(19) World Intellectual Property Organization

International Bureau



A CORRECT CONTROL OF CORRECT CONTROL OF CORRECT CORREC

(43) International Publication Date 29 July 2004 (29.07.2004)

PCT

(10) International Publication Number WO 2004/062338 A3

(51) International Patent Classification⁷:

C12Q 1/68

(21) International Application Number:

PCT/GB2004/000127

(22) International Filing Date: 14 January 2004 (14.01.2004)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data: 0300802.6

14 January 2003 (14.01.2003) GI

(71) Applicants and

- (72) Inventors: TISI, Lawrence, C. [GB/GB]; 116 Lynn Road, Ely CB6 1DE (GB). MURRAY, James, A., H. [GB/GB]; 4 Benson Street, Cambridge CB2 1TN (GB).
- (74) Agents: HALLYBONE, Huw, George et al.; Carpmaels & Ransford, 43-45 Bloomsbury Square, London WC1A 2RA (GB).
- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI,

GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments
- (88) Date of publication of the international search report:

 10 September 2004

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: METHOD FOR DETERMINING THE AMOUNT OF TEMPLATE NUCLEIC ACID PRESENT IN A SAMPLE

(57) Abstract: A method for determining the amount of template nucleic acid present in a sample comprising the steps of: i) bringing into association with the sample all the components necessary for nucleic acid amplification, and all the components necessary for a bioluminescence assay for nucleic acid amplification and subsequently: ii) performing the nucleic acid amplification, iii) monitoring the intensity of light output from the bioluminescence assay; and iv) determining the amount of template nucleic acid present in the sample.

	INTERNATIONAL SEARCH REP	ORT	International Ap	
A. CLASS	SECATION OF SUBJECT MATTER		PCT/GB200	04/000127
IPC 7	SIFICATION OF SUBJECT MATTER C12Q1/68			-
According	to International Patent Classification (IPC) or to both national classific	ation and IPC		
	SEARCHED			
IPC 7	ocumentation searched (classification system followed by classification ${\tt C12Q}$	ion symbols)		
Documents	tion searched other than minimum documentation to the extent that s	such documents are inclu	ided in the fields s	earched
	data base consulted during the international search (name of data ba	se and, where practical,	search terms used	J)
EPO-In	ternal, WPI Data, EMBASE, BIOSIS			
	ENTS CONSIDERED TO BE RELEVANT			
Category *	Citation of document, with Indication, where appropriate, of the rele	evant passages		Relevant to claim No.
X	WO 01/42496 A (DZIEGLEWSKA HANNA ;PYROSEQUENCING AB (SE); LUNDEBER (SE);) 14 June 2001 (2001-06-14)	G JOAKIM		1-7,14, 15, 22-25, 28,29, 32,34,
	page 14, line 17 - page 16, line 43, line 30 - page 44, line 33; f 6-7; claims 1-11, 22-23	6; page igures		36,37
	_	/		
[V] Surb	Or documents and K. L. H. H.			
	er documents are listed in the continuation of box C.	X Patent family me	embers are listed in	annex.
 'L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) 'O' document referring to an oral disclosure, use, exhibition or other means 'P' document published prior to the international filing date but later than the priority date claimed 		 "T' later document published after the international fliing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family 		
	clual completion of the international search	Date of mailing of the	International searc	ch report
	June 2004	19/07/20	04	
mame and ma	alling address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo ni,	Authorized officer		
	Fax: (+31-70) 340-3016	Hennard,	С	

Form PCT/ISA/210 (second sheet) (January 2004)

International Application No
PCT/GB2004/000127

Category °	NYGREN M ET AL: "QUANTIFICATION OF HIV-1 USING MULTIPLE QUANTITATIVE POLYMERASE CHAIN REACTION STANDARDS AND	1-7,14, 15,
Х	CHAIN REACTION STANDARDS AND	15,
	BIOLUMINOMETRIC DETECTION" ANALYTICAL BIOCHEMISTRY, ACADEMIC PRESS, NEW YORK, NY, US, vol. 288, no. 1, 1 January 2001 (2001-01-01), pages 28-38, XP000977141 ISSN: 0003-2697 page 30, right-hand column; page 32, left-hand column, lines 15-18	22-25, 28,29, 32,34, 36,37
<i>(</i>	WO 02/064830 A (MURRAY JAMES AUGUSTUS HENRY; TISI LAURENCE CARLO (GB); UNIV CAMBRI) 22 August 2002 (2002-08-22) claim 17; examples	34-36
'	US 6 258 568 B1 (NYREN PAL) 10 July 2001 (2001-07-10) claim 16	34-36
	MORI YASUYOSHI ET AL: "Detection of loop-mediated isothermal amplification reaction by turbidity derived from magnesium pyrophosphate formation" BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, ACADEMIC PRESS, SAN DIEGO, CA, US, vol. 289, no. 1, 23 November 2001 (2001-11-23), pages 150-154, XP002255232 ISSN: 0006-291X cited in the application abstract; page 153, left-hand column, last paragraph; page 154, forlast paragraph	1-37
	TABARY T ET AL: "HOMOGENEOUS PHASE PYROPHOSPHATE (PPI) MEASUREMENT (H3PIM). \A NON-RADIOACTIVE, QUANTITATIVE DETECTION SYSTEM FOR NUCLEIC ACID SPECIFIC HYBRIDIZATION METHODOLOGIES INCLUDING GENE AMPLIFICATION" JOURNAL OF IMMUNOLOGICAL METHODS, ELSEVIER SCIENCE PUBLISHERS B.V., AMSTERDAM, NL, vol. 156, no. 1, 1992, pages 55-60, XP000320297 ISSN: 0022-1759 cited in the application abstract	1-37
	-/	1

International Application No
PCT/GB2004/000127

C.(Continueti	ion) DOCUMENTS CONSIDERED TO BE RELEVANT	PCT/GB2004/000127	
	Citation of document, with Indication, where appropriate, of the relevant passages	Relevant to claim No.	
A	WHITE ET AL: "improved thermostability of the north american firefly luciferase: saturation mutagenesis at position 354" BIOCHEMICAL JOURNAL, PORTLAND PRESS, LONDON, GB, vol. 319, 1996, pages 343-350, XP002097112 ISSN: 0264-6021 introduction, page 343	1-37	
4	HIGUCHI R ET AL: "KINETIC PCR ANALYSIS: REAL—TIME MONITORING OF DNA AMPLIFICATION REACTIONS" BIO/TECHNOLOGY, NATURE PUBLISHING CO. NEW YORK, US, vol. 11, no. 9, 1 September 1993 (1993—09—01), pages 1026—1030, XP000197685 ISSN: 0733—222X the whole document	1-37	

International application No.

PCT/GB2004/000127

Вох	No. I	Nucleotide and/or amino acid sequence(s) (Continuation of item 1.b of the first sheet)
1.	With	regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed not on the basis of:
	a.	type of material
		X a sequence listing
		table(s) related to the sequence listing
	b.	format of material
		X In written format
		X In computer readable form
	C.	time of filing/fumishing
		contained in the international application as filed
		filed together with the international application in computer readable form
		X furnished subsequently to this Authority for the purpose of search
2.		In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished.
•	A _4 _0*a*	
3.	Additi	ional comments:

information on patent family members

International Application No PCT/GB2004/000127

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 0142496	A	14-06-2001	AU WO US	2192801 A 0142496 A2 2003157499 A1	18-06-2001 14-06-2001 21-08-2003
WO 02064830	A 	22-08-2002	CA EP WO	2440698 A1 1360330 A2 02064830 A2	22-08-2002 12-11-2003 22-08-2002
US 6258568	B1	10-07-2001	AU DE DE EP WO JP US	5331198 A 69713386 D1 69713386 T2 0946752 A1 9828440 A1 2001506864 T 2003104372 A1	17-07-1998 18-07-2002 30-01-2003 06-10-1999 02-07-1998 29-05-2001 05-06-2003